

2. (Twice Amended) [A] An isolated polypeptide characterized in that it: [-]

- (a) is free of botulinum toxin activity,
- (b) is free of toxoid, and
- (c) is capable of eliciting, in a mammal, an immunological response that is protective against type F botulinum toxin.

3. (Twice Amended) [A] An isolated polypeptide according to Claim 2 comprising a fragment or a derivative of a heavy chain of a type F botulinum neurotoxin.

4. (Twice Amended) [A] An isolated polypeptide according to Claim 3 wherein said fragment or said derivative is up to 600 amino acids long.

5. (Twice Amended) [A] An isolated polypeptide according to Claim 3 wherein said fragment is selected from the group consisting of:

- (a) amino acids 848-1278 of a type F botulinum toxin, *product*
- (b) amino acids 848-991 of a type F botulinum toxin,
- (c) amino acids 992-1135 of a type F botulinum toxin, and
- (d) amino acids 1136-1278 of a type F botulinum toxin.

6. (Twice Amended) [A] An isolated polypeptide according to Claim 3 wherein said derivative comprises a dimer of the fragment [according to any of (a)-(d) of Claim 5] selected from the group consisting of: *dimer*

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- (a) amino acids 848-1278 of a type F botulinum toxin,  
(b) amino acids 848-991 of a type F botulinum toxin,  
(c) amino acids 992-1135 of a type F botulinum toxin, and  
(d) amino acids 1136-1278 of a type F botulinum toxin.

6 1 *Salva*  
(Twice Amended) A polypeptide composition for use in manufacture of a vaccine,  
said composition comprising: [-]

- (1) an isolated [a] polypeptide free of toxin activity and capable of inducing, in a  
mammal, protective immunity against a botulinum toxin; and  
(2) a polypeptide [adapted to facilitate or enhance] that facilitates or enhances  
purification of the composition.

6 2 *Salva*  
(Twice Amended) A polypeptide composition according to Claim 7 comprising: [-]  
(1) [a] an isolated polypeptide according to Claim 2; and  
(2) [a] an isolated polypeptide [adapted to bind to] that binds a chromatography  
column.

6 7  
10. (Twice Amended) A polypeptide composition according to Claim 7 comprising an  
isolated polypeptide [adapted to bind to] that binds an affinity chromatography column.

6 3 *Salva*  
17. (Twice Amended) A method of making a pharmaceutical composition comprising: [-]  
(a) expressing in a host cell a DNA encoding a fusion protein, said protein being a  
fusion of (i) a polypeptide free of toxin activity and capable of inducing protective immunity